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Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number. Substitute for form 1449A/PTO Complete if Known 10/526,893 Application Number INFORMATION DISCLOSURE Herewith Filing Date STATEMENT BY APPLICANT First Named Inventor David E. Lowery Art Unit To Be Determined (Use as many sheets as necessary) Examiner Name To Be Determined PHRM0002-105 Sheet of 19 Attorney Docket Number

			U.S. PATENT D	OCUMENTS	
Examiner	Cite	Document Number	Publication/Issue Date	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant
Initials *	No.1	Number - Kind Code <sup>2</sup> (if known)	MM-DD-YYYY	Cited Document	Passages or Relevant Figures Appear
	AA	US-4,343,940	8-10-1982	Kreighbaum et al.	
	AB	US-4,399,216	8-16-1983	Axel et al.	
	AC	US-4,447,608	5-08-1984	Jones et al.	
	AD	US-4,683,195	7-28-1987	Mullis et al.	
	AE	US-4,683,202	7-28-1987	Mullis	
	AF	US-4,757,072	7-12-1988	Kabbe et al.	
	AG	US-4,879,236	11-07-1989	Smith et al.	
	AH	US-5,217,999	6-08-1993	Levitzki et al.	
	ΑI	US-5,302,606	4-12-1994	Spada et al.	
	AH	US-5,316,553	5-31-1994	Kaul et al.	
	AK	US-5,330,992	7-19-1994	Eissenstat et al.	
	AL	US-5,585,277	12-17-1996	Bowie et al.	
	AM	US-5,753,615	5-19-1998	Thorpe et al.	
	AN	US-5,880,141	3-09-1999	Tang et al.	
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		FOREIGN PA	TENT DOCUM	MENTS		
	Cita	Foreign Patent Document	Publication	Name of Patentee or	Pages, Columns, Lines,	
Examiner Initials*	Cite No. <sup>1</sup>	Country Code <sup>3</sup> - Number <sup>4</sup> - Kind Code <sup>5</sup> (if known)	Date/Filing Date MM-DD-YYYY	Applicant of Cited Document	where Relevant Passages or Relevant Figures Appear  ex Corp al Chem Ltd. et al. s et al	T <sup>6</sup>
	AO	EP 0 367 566	05-1990	Immunex Corp		
	AP	EP 0 520 722	12-1992	Imperial Chem		
	AQ	EP 0 562 734	09-1993	Zeneca Ltd. et al.		
	AR	WO01/70981	9/27/01	Ebens et al		
	AS	WO 91/09955	07-1991	App. Research		
	AT	WO 91/15495	10-1991	Pfizer		
	AU	WO 91/18982	12-1991	Immunex Corp		
	AV	WO 92/20642	11-1992	Rhone-Poulenc		
	AW	WO 92/20808	11-1992	Cell Genesys		

Examiner Signature	Date Considered	

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Substitu	ute for form 1449/	VPTO			Complete if Known	
				Application Number	10/526,893	
INF	ORMATIC	ON DIS	CLOSURE	Filing Date	Herewith	
STA	TEMENT	BY A	PPLICANT	First Named Inventor	David E. Lowery	
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	(Use as man	y sheets as	necessary)	Examiner Name	To Be Determined	
Sheet	2	of	19	Attorney Docket Number	PHRM0002-105	

	U.S. PATENT DOCUMENTS						
Examiner	Cite	Document Number	Publication/Issue Date	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant		
Initials *	No.1	Number - Kind Code <sup>2</sup> (if known)	MM-DD-YYYY	Original Description	Passages or Relevant Figures Appear		
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		FOREIGN PA	TENT DOCUM	MENTS		
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	AX	WO 92/21660	12-1992	Pfizer		
	AY	WO 93/11236	06-1993	Med. Research Council		
	AZ	WO 94/03427	02-1994	Warner-Lambert		
	BA	WO 94/12650	06-1994	Transkaryotic		
	ВВ	WO 94/14808	07-1994	Farmitalia Carlo Erba S.R.L.		
	BC	WO 95/20652	08-1995	Medigene		
	BD	WO 96/22976	08-1996	Pharmacia SPA		
	BE	WO 97/09433	03-1997	Med. Research Counsil		
	BF	WO 98/37177	08-1998	MS State Univ.		
	IE	WO 01/71042	09-27-2001	PE Corp.		
	ĬF	WO01/70980	09/27/01	Cravchik		
	IG	EP 0 566 266	10-1993			

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Complete if Known Substitute for form 1449B/PTO **Application Number** 10/526,893 INFORMATION DISCLOSURE Filing Date Herewith STATEMENT BY APPLICANT First Named Inventor David E. Lowery Art Unit To Be Determined (Use as many sheets as necessary) Examiner Name To Be Determined Attorney Docket Number PHRM0002-105 Sheet

		NON PATENT LITERATURE DOCUMENTS	
Examiner Initials *	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T²
	ВG	Allen et al., "Modulation of CD4 by Suramin", Clin. Exp. Immunol., 1993, vol. 91, pp. 141-156.	
· · · · · ·	ВН	Altschul et al., "Gapped BLAST and PSI-BLAST: a new generation of protein database search programs," Nucl. Acids Res., 1997, 25(17), 3389-3402.	
	ВІ	Altschul et al., "Basic Local Alignment Search Tool," J. Mol. Biol., 1990, 215, 403-410.	
	ВЈ	Anafi et al., "Tyrphostin-Induced Inhibition of p210.sup.bcr-abl Tyrosine Kinase Activity Induces K562 to Differentiate", Blood, 1993, vol. 82, No. 12, pp. 3524-3529.	
	BK	Anderson, W. F., "Human gene therapy," Science, 1992, 256, 808-813.	
	BL	Aukrust et al., "Enhanced Levels of Soluble and Membrane-Bound CD40 Ligand in Patients with Unstable Angina. Possible Reflection of T Lymphocyte and Platelet Involvement in the Pathogenesis of Acute Coronary Syndromes", Circulation, 1999, vol. 100, pp. 614-620.	
	ВМ	Ausubel, et al. (Eds.), "Chapter 6, Screening of recombinant DNA libraries," Current Protocols in Molecular Biology, 1994, John Wiley & Sons, 6.0.1-6.4.10.	
	BN	Baindur et al., "Selective fluorescent ligands for pharmacological receptors," Drug Dev. Res., 1994, 33, 373-398.	
	во	Baker et al., "Induction of Acetylcholine Receptor Clustering by Native Polystyrene Beads. Implication of an Endogenous Muscle-derived Signalling System", J. Cell. Sci., 1992, vol. 102, pp. 543-555.	
	ВР	Barker et al., "In-Vitro Activity of Non-glutamate Containing Quinazoline-based Thymidylate Synthase Inhibitors", Proc. of Am. Assoc. for Cancer Res., 1991, vol. 32, p. 327.	
	BQ	Benoist et al., "In vivo sequence requirements of the SV40 early promoter region," Nature, 1981, 290, 304-310.	

Examiner	Date	
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<sup>\*</sup>EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance

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		NON PATENT LITERATURE DOCUMENTS	·
Examiner Initials *	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T²
	BR	Bertino, Cancer Res., "Toward Improved Selectivity in Cancer Chemotherapy: The Richard and Hinda Rosenthal Foundation Award Lecture", 1979, vol. 3, pp. 293-304.	
	BS	Bilder et al., "Tyrphostins Inhibit PDGF-induced DNA Synthesis and Associated Early Events in Smooth Muscle Cells", Amer. Physiol. Soc., 1991, pp. 6363-6143:C721-C730.	
	вт	Birgul, N. et al., "Reverse physiology in Drosophila: Identification of a novel allatostatin-like neuropeptide and its cognate receptor structurally related to the mammalian somatostatin/galanin/opioid receptor family", The EMBO Journal, 1999, 18(21), 5892-5900.	
	BU	Bohm, S. K., et al., "Regulatory mechanisms that modulate signalling by G-protein-coupled receptors," Biochem. J., 1997, 322, 1-18.	
	BV	Bosse, R., et al., "Development of nonseparation binding and functional assays for G protein-coupled receptors for high throughput screening: Pharmacological characterization of the immobilized CCR5 receptor on FlashPlate.RTM.," J. Biomolecular Screening, 1998, 3(4), 285-292.	
	BW	Boulton, T. G., et al., "ERKs: A family of protein-serine/threonine kinases that are activated and tyrosine phosphorylated in response to insulin and NGF," Cell, 1991, 65, 663-675.	
	ВХ	Brunton, V. G., et al., Proceedings of Amer. Assoc. Cancer Res., No. 3335, 1992, 33, 558.	
	BY	Bryckaert, M., et al., "Inhibition of platelet-derived growth factor-induced mitogenesis and tyrosine kinase activity in cultured bond marrow fibroblasts by tyrphostins," Experimental Cell Research, 1992, 199, 255-261.	
	BZ	Burke, T. R., et al., "Bicyclic compounds as ring-constrained inhibitors of protein-tyrosine kinase p56.sup.ick," J. Med. Chem., 1993, 36(4), 425-432.	
	CA	Burke, T. R., et al., "Arylamides of hydroxylated isoquinolines as protein-tyrosine kinaseinhibitors,"BioOrganic Med. Chem. Ltrs., 1992, 2(12), 1771-1774.	
	СВ	Capecchi, M. R., "Altering the genome by homologous recombination," Science, 1989, 244, 1288-1292.	

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Signature	Considered	

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	СС	Chambers, R. C., et al., "Thrombin stimulates fibroblast procollagen production via proteolytic activation of protease-activated receptor 1," Biochem J., 1998, 333, 121-127.	
	CD	Choo, Y., et al., "Promoter-specific activation of gene expression directed by bacteriophage-selected zinc fingers," J. Mol. Biol., 1997, 273, 525-532.	
	CE	Cicala, C., et al., "Bronchoconstrictor effect of thrombin and thrombin receptor activating peptide in guinea-pigs in vivo," Br. J. Pharmacol, 1999, 126, 478-484.	
	CF	Cirino, G., et al., "Thrombin functions as an inflammatory mediator through activation of its receptor," J. Exp. Med., 1996, 183, 821-827.	
	CG	Colotta, F., et al., "Expression of monocyte chemotactic protein-1 by monocytes and endothelial cells exposed to thrombin," Am. J. Pathol, 1994, 144, 975-985.	
	СН	Cosman, D., et al., "High Level Stable Expression of Human Interleukin-2 receptors in Mouse Cells Generates only Low Affinity Interleukin-2 Binding Sites," Mol. Immunol., 1986, 23(9), 935-941.	
	CI	Cosman, D., et al., "Cloning, sequence and expression of human interleukin-2 receptor," Nature, 1984, 312, 768-771.	
	CJ	Curtin, N. J., et al., "Inhibition of the growth of human hepatocellular carcinoma in vitro and in athymic mice by a quinazoline inhibitor of thymidylate synthase, CB3717," J. Cancer, 1986, 53, 361-368.	
	СК	Dayoff, in Atlas of Protein Sequence and Structure, 1972, National Biochemical Research Foundation, Washington, D.C., 5, 124.	
	CL	DiCuccio, M. N., et al., "A functional tethered ligand thrombin receptor is present on human hematopoietic progenitor cells," Exp. Hematol, 1996, 24, 914-918.	
-	СМ	Dolle, R. E., et al., "5,7-dimethoxy-3-(4-pyridinyl)quinoline is a potent and selective inhibitor of human vascular .betatype platelet-derived growth factor receptor tyrosine kinase," J. Med. Chem., 1994, 37, 2627-2629.	

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Signature	Considered	

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	CN	Dong, Z., et al., "Activation of tumoricidal properties in macrophages by lipopolysaccharide requirements protein-tyrosine kinase activity," J. Leukocyte Biology, 1993, 53, 53-60.	
	со	Dong, Z., et al., "Protein tyrosine kinase inhibitors decrease induction of nitric oxide synthase activity in lipopolysaccharide-responsive and lipopolysaccharide-nonresponsive murine macrophages," J. Immunol., 1993, 151(5), 2717-2724.	
	СР	Donovan, F. M., et al., "Thrombin induces apoptosis in cultured neurons and astrocytes via a pathway requiring tyrosin kinase and RhoA activities," J. Neurosci., 1997, 17(14), 5316-5326.	
	CQ	Dooley, C. T., et al., "Binding and in vitro activities of peptides with high affinity for the nociceptin/orphanin FQ receptor, ORL1," J. Pharmacology and Experimental Therapeutics, 1997, 283(2), 735-741.	
	CR	Dunlop, J., et al., "Characterization of 5-HT.sub.1A receptor functional coupling in cells expressing the human 5-HT.sub.1A receptor as assessed with the cytosensor microphysiometer," J. Pharmacological and Toxicological Methods, 1998, 40(1), 47-55.	
	CS	Fernandes, D. J., et al., "Biochemical and antitumor effects of 5,8-dideazaisopteroylglutamate, a unique quinazoline inhibitor of thymidylate synthase," Cancer Research, 1983, 43, 1117-1123.	
	СТ	Ferris, J. P., et al., "Synthesis of Quinazoline Nucleosides from Ribose and Anthranilonitrile. Application of Phase-Transfer Catalysis in Nucleoside Synthesis," J. Org. Chem., 1979, 44(2), 173-178.	
	CU	Fields, S., et al., "A novel genetic system to detect protein-protein interactions," Nature, 1989, 340, 245-246.	
	CV	Fields, S., et al., "The two-hybrid system: an assay for protein-protein interactions," Trends in Genetics, 1994, 10, 286-292.	
	CW	Foote, J., et al., Antibody framework residues affecting the conformation of the hypervariable loops, J. Mol. Biol., 1992, 224, 487-499.	
	СХ	Frandsen, E. K., et al., "A simple ultrasensitive method for the assay of cyclic AMP and CMP in tissues," Life Sciences, 1976, 529-542.	

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	CY	Fry, D.W., et al., "A specific inhibitor of the epidermal growth factor receptor tyrosine kinase," Science, 1994, 265, 1093-1095.	
	cz	Gazit, A., et al., "Tyrphostins I: Synthesis and biological activity of protein tyrosine kinase inhibitors," J. Med. Chem., 1989, 32, 2344-2352.	
	DA	Gazit, A., et al., "Tyrphostins. 3. Structure-activity relationship studies of .alphasubstituted benzylidenemalononitrile 5-S-aryltyrphostins," J. Med. Chem., 1993, 36, 3556-3564.	
	DB	George, S. E., et al., "Evaluation of a CRE-directed luciferase reporter gene assay as an alternative to measuring cAMP accumulation," J. Biomolecular Screening, 1997, 2(4), 235-240.	
	DC	Gerhardt, C. C., et al., "Functional characteristics of heterologously expressed 5-HT receptors," Eur. J. Pharmacology, 1997, 334, 1-23.	
	DD	Gill, J. S., et al., "Thrombin induced inhibition of neurite outgrowth from dorsal root ganglion neurons," Brain Res., 1998, 797, 321-327.	
	DE	Grabham, P., et al., Thrombin receptor activation stimulates astrocyte proliferation and reversal of stellation by distinct pathways: involvement of tyrosine phosphorylation, J. Neurochem, 1995, 64, 583-591.	
	DF	Greisman, H. A., et al., "A general strategy for selecting high-affinity zinc finger proteins for diverse DNA target sites," Science, 1997, 275, 657-661.	
	DG	Guerrero, F. D., "Transcriptional Expression of a Putative Tachykinin-like Peptide Receptor Gene From Stable Fly.sup.1," Peptides, 1997, 18(1), 1-5.	
	DH	Hauck, R. W., et al., ".alphathrombin stimulates contraction of human bronchial rings by activation of protease-activated receptors," Am J. Physiol, 1999, 277, L22-L29.	
	DI	Hauser, F., et al., "Molecular Cloning, Genomic Organization, and Developmental Regulation of a Novel Receptor from Drosophila melanogaster Structurally Related to Members of the Thyroid-stimulating Hormone, Follicle-stimulating Hormone, Luteinizing Hormone/ Choriogonadotropin Receptor Family from Mammals," The J. of Biological Chemistry, 1997, 272(2), 1002-1010.	

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<sup>\*</sup>EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

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Substitute for form 1449B/PTO	<u></u>	Complete if Known	
INFORMATION DIGGLOG	Application Number	10/526,893	
INFORMATION DISCLOS	rilling Date	Herewith	
STATEMENT BY APPLIC	ANT First Named Inventor	David E. Lowery	
	Art Unit	To Be Determined	
(Use as many sheets as necessary	Examiner Name	To Be Determined	
Sheet 8 of 19	Attorney Docket Numb	ber PHRM0002-105	

		NON PATENT LITERATURE DOCUMENTS	<del></del>
Examiner Initials *	Cite No.1	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T²
	DJ	Hauser, F. et al., "Molecular Cloning, Genomic Organization, and Developmental Regulation of a Novel Receptor from Drosophila melanogaster Structurally Related to Gonadotropin-Releasing Hormone Receptors from Vertebrates," Biochem. Biophys. Res. Comm., 1998, 249, 822-828.	
	DK	Henikoff, S., et al., "Amino acid substitution matrices from protein blocks," Proc. Natl. Acad. Sci. USA, 1992, 89, 10915-10919.	
	DL	Hill, D. C., "Trends in development of high-throughput screening technologies for rapid discovery of novel drugs," Cur. Opinion Drug Disc. Dev., 1998, 1(1), 92-97.	
	DM	Hodgson, J., "Receptor screening and the search for new pharmacteuticals," Bio/Technology, 1992, 10, 973-980.	
	DN	Hoffman, M., et al., "Thrombin enhances monocyte secretion of tumor necrosis factor and interleukin-1 beta by two distinct mechanisms," Blood Cells Mol Dis, 1995, 21, 156-167.	
	DO	Jackman, A. L., et al., "ICID1694, a quinazoline antifolate thymidylate synthase inhibitor that is a potent inhibitor of L1210 tumor cell growth in vitro and in vivo: A new agent for clinical study," Cancer Research, 1981, 51, 5579-5586.	
-	DP	Jayawickreme, C. K., et al., Gene expression systems in the development of high-throughput screens, Current Opinion in Biotechnology, 1997, 8, 629-634.	
	DQ	Jones, P. T., et al., "Replacing the compementarity-determining regions in a human antibody with those from a mouse," Nature, 1986, 321, 522-525.	
	DR	Jones, T. R., et al., "Quinazoline Antifolates Inihibiting Thymidylate Synthase: Variation of the Amino Acid," J. Med Chem., 1986, 29, 1114-1118.	
	DS	Kanterman, R. Y., et al., "Transfected D.sub.2 dopamine receptors mediate the potentiation of arachidonic acid release in chinese hamster ovary cells," Molecular Pharmacology, 1991, 39, 364-369.	
	DT	Karlin, S., et al., "Applications and statistics for multiple high-scoring segments in molecular sequences," Proc. Natl. Acad. Sci. USA, 1993, 90, 5873-5787.	

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	DU	Kaur, G., et al., "Tyrphostin induced growth inhibition: corelation with effect on p210.sup.berabl autokinase activity in K562 chronic myelogenous leukemia," Anti-Cancer Drugs, 1994, 5, 213-222.	
	DV	Kettleborough, C. A., et al., "Humanization of a mouse monoclonal antibody by CDR-grafting: the importance of framework residues on loop conformation," Protein Engin., 1991, 4(7), 773-783.	•
	DW	Kim, J., et al., "Design of TATA box-binding protein/zinc finger fusions for targeted regulation of gene expression," Proc. Natl. Acad. Sci. USA, 1997, 94, 3616-3620.	
	DX	King, M. J., et al., "Site-specific dephosphorylation and deactivation of the human insulin receptor tyrosine kinase by particulate and soluble phosphotyrosyl protein phosphatases," Biochem. J., 1991, 275, 413-418.	
	DY	Kowal, D., et al., "A [.sup.35 S]GTP.gamma.S binding assessment of metabotropic glutamate receptor standards in chinese hamster ovary cell lines expressing the human metabotropic receptor subtypes 2 and 4," Neuropharmacology, 1998, 37, 179-187.	
	DZ	Kuntzweiler, T. A., et al., "Rapid assessment of ligand actions with nicotinic acetylcholine receptors using calcium dynamics and FLIPR," Drug Development Research, 1998, 44(1), 14-20.	
	EA	Kuo, M., et al., "Effects of signalling transduction modulators on the transformed phenotypes in v-H-ras-transformed NIH 3T3 cells," Cancer Letters, 1993, 74, 197-202.	
-	ЕВ	Lajiness et al., "D2 dopamine receptor stimulation of mitogenesis in transfected chinese hamster ovary cells: relationship to dopamine stimulation of tyrosine phosphorylations", J. Pharm. Exp. Ther., 1993, vol. 267, No. 3, 1573-1581.	
	EC	Lee, C., et al., "Active-site directed reductive alkylation of xanthine oxidase by imidazo[4,5-g]quinazoline-4,9-diones functionalized with a leaving group," Biochemistry, 1987, 26(23), 7355-7362.	
	ED	Lehninger, "Chapter 4, The amino acid building blocks of proteins," Biochemistry, 2.sup.nd Ed., 1975, Worth Publishers, Inc., New York, New York, 71-77.	
	EE	Lemus, et al., "Studies of extended quinone methides. Synthesis and physical studies of purine-like monofunctional and bifunctional imidazo[4,5-g]quinazoline reductive alkylating agents," J. Org. Chem., 1989, 54, 3611-3618.	

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	EF	Lenz, C. et al., "Molecular Cloning and Genomic Organization of a Second Probable Allastatin Receptor from Drosophila melanogaster", Biochem. Biophys. Res. Comm., 2000, 273, 571-577.	
	EG	Lenz, C. et al., Drosophila melanogaster allatostatin G-protein receptor mRNA, complete cds, GenBank Accession No. AF253526, Jul. 14, 2000.	
	EH	Lenz, C. et al., "Molecular Cloning and Genomic Organization of an Allatostatin Preprohormone from Drosophila melanogaster", Biochem. Biophys. Res. Comm., 2000, 273, 1126-1131.	
	EI	Levitzki, A., "Tyrphostins: tyrosine kinase blockers as novel antiproliferative agents and dissectors of signal transduction," The FASEB J., 1992, 6, 3275-3282.	
	EJ	Ley, K., et al., "Synthesen unter verwendung von benzofuroxan," Synthesis, 1975, 415-522 (English abstract).	
	EK	Li, X-J., et al., "Cloning, heterologous expression and developmental regulation of a Drosophila receptor for tachykinin-like peptides," The EMBO Journal, 1991, 10(11), 3221-3229.	
	EL	Li, X-J., et al., "Cloning, Functional Expression, and Developmental Regulation of a Neuropeptide Y Receptor from Drosophila melanogaster," The J. of Biological Chemistry, 1992, 267(1), 9-12.	
	EM	Li, XJ. et al., D. melanogaster neuropeptide receptor mRNA, complete cds, GenBank Accession No. M81490, Apr. 26, 1993.	
	EN	Lin, A. H., et al., "The oxazolidinone eperezolid binds to the 50S ribosomal subunit and competes with binding of chloramphenicol and lincomycin," Antimicrobial Agents and Chemotherapy, 1997, 41(10), 2127-2131.	
	EO	Liu, Q., et al., "Design of polydactyl zinc-finger proteins for unique addressing within complex genomes," Proc. Natl. Acad. Sci. USA, 1997, 94, 5525-5530.	
	EP	Luckow, V. A., et al., "High Level Expression of Nonfused Foreign Genes with Autographa californica Nuclear Polyhedrosis Virus Expression Vectors," Virology, 1989, 170, 31-39.	

Signature Considered Considered	Examiner Signature	Date Considered
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INFORMATION DISCLOSURE	Filing Date	Herewith		
STATEMENT BY APPLICANT	First Named Inventor	David E. Lowery		
	Art Unit	To Be Determined		
(Use as many sheets as necessary)	Examiner Name	To Be Determined		
Sheet 11 of 19	Attorney Docket Number	PHRM0002-105		

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	EQ	Luckow, V. A., et al., "Trends in the development of baculovirus expression vectors," Bio/Technology, 1988, 6, 47-55.	
	ER	Lyall, R. M., et al., "Tyrphostins inhibit epidermal growth factor (EGF)-receptor tyrosine kinase activity in living cells and EGF-stimulated cell preliferation," J. Biol. Chem., 1989, 264, 14503-14509.	
-	ES	Maguire, M. P., et al., "A new series of PDGF receptor tyrosine kinase inhibitors: 3-substituted quinoline derivatives," J. Med. Chem., 1994, 37, 2129-2131.	
	ET	Maxwell, R. J., et al., ".sup.19 F nuclear magnetic resonance imaging of drug distribution in vivo: The disposition of an antifolate anticancer drug in mice," Magnetic Resonance in Medicine, 1991, 17, 189-196.	
	EU	McColl, D. J., et al., "Structure-based design of an RNA-binding zinc finger", Proc. Natl. Acad. Sci. (USA), 1997, vol. 96, 9521-9526.	
	EV	Mini, E., et al., "Cytotoxic effects of folate antagonists against methotrexate-resistant human leukemic lymphoblast CCRF-CEM cell lines," Cancer Res., 1985, 45, 325-330.	
	EW	Monnier, D., et al., "NKD, a Developmentally Regulated Tachykinin Receptor in Drosophila," The J. of Biological Chemistry, 1992, 267(2), 1298-1302.	
	EX	Monnier, D. et al., Drosophila melanogaster tachykinin receptor (NKD) mRNA, complete cds, GenBank Accession No. M77168, Apr. 26, 1993.	
	EY	Morris, R., et al., "Thrombin receptor expression in rheumatoid and osteoarthritic synovial tissue", Ann. Rheum. Dis., 1996, vol. 55, 841-843.	
	EZ	Morrison, et al., "Genetically engineered antibody molecules," Dixon, F.J., et al. (Eds.), Adv. Immunol., 1989, 44, 65-92.	
_	FA	Murphy, A. J., et al., "From DNA to drugs: the orphan G-protein coupled receptors," Cur. Opinion Drug Disc. Dev., 1998, 1(2), 192-199.	

Examiner	Date	
Signature	Considered	

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	FB	Myers, P., "Will combinatorial chemistry deliver real medicines," Curr. Opin. Biotechnology, 1997, 8, 701-707.	
	FC	Nachman & Homan, in Insect Neuropeptides; Chemistry, Biology and Action, Menn, Kelly & Massler, Eds., 1991, 194-214, American Chemical Society, Washington, DC.	
	FD	Nakayama, G. R., "Microplate assays for high-throughput screening," Cur. Opinion Drug Disc. Dev., 1998, 1, 85-91.	
	FE	Naldini, A., et al., "Thrombin modulation of natural killer activity in human peripheral lymphocytes," Cell Immunol, 1996, 172, 35-42.	
	FF	Nambu et al., "Isolation and Characterization of a Drosophila Neuropeptide Gene", Neuron, 1988, 1, 55-61.	
	FG	Nichols, R. et al., "Identification and Characterization of a Drosophila Homologue to the Vertebrate Neuropeptide Cholecystokinin", J. Biol. Chem., 1988, 263, 12167-12170.	
	FH	Okayama, H., et al., "A cDNA cloning vector that permits expression of cDNA inserts in mammalian cells," Mol. Cell. Biol., 1983, 3(2), 280-289.	
	FI	Padlan, E. A., "A possible procedure for reducing the immunogenicity of antibody variable domains while preserving their ligand-binding properties," Molecular Immunol., 1991, 28(4/5), 489-498.	
	FJ	Pausch, M. H., "G-protein-coupled receptors in Saccharomyces cerevisiae: high-throughput screening assays for drug discovery," Trends in Biotechnology, 1997, 15, 487-494.	
_	FK	Peterson, G., et al., "Genistein and biochanin A inhibit the growth of human prostate cancer cells but not epidermal growth factor receptor tyrosine autophosphorylation," The Prostate, 1993, 22, 335-345.	
	FL	Phillips, S. D., et al., "Quino[1,2-c]quinazolines. I. Synthesis of quino[1,2-c]quinazolinium derivatives and the related indazolo[2,3-a]quinoline derivatives as analogs of the antitumor benzol[c]phenanthridine alkaloids," J. Heterocyclic Chem., 1980, 17(19), 1489-1596.	

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STAT	EMENT B	BY A	PPLICANT	First Named Inventor	David E. Lowery
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	FM	Pillemer, G., et al., "Insulin dependence of murine lymphoid T-cell leukemia," Int. J. Cancer, 1992, 50, 80-85.	
	FN	Pindon, A., et al., "Thrombin-induced reversal if astricyte stellation is mediated by activation of protein kinase C .beta1," Eur. J. Biochem., 1998, 255, 766-774.	
	FO	Posner, I., et al., "Kinetics of inhibition by tyrophostins of the tyrosine kinase activity of the epidermal growth factor receptor and analysis," Molecular Pharmacology, 1993, 45, 673-683.	
_	FP	Reece, P. A., et al., "Pharmacokinetics of trimetrexate administered by five-day continuous infusion to patients with advanced cancer," Cancer Research, 1977, 47(11), 2996-2999.	
	FQ	Rendu, F., et al., "Inhibition of platelet activation by tyrosine kinase inhibitors," Biol. Pharmacology, 1992, 44(5), 881-888.	
	FR	Riechmann, L., et al., "Reshaping human antibodies for therapy," Nature, 1988, 332, 323-327.	
	FS	Rogers, M. V., "Light on high-throughput screening: fluorescence-based assay technologies," Drug Discovery Today, 1997, 2(4), 156-160.	
	FT	Sauro, M. D., et al., "Tyrphostin attenuates platelet-derived growth factor-induced contraction in aortic smooth muscle through inhibition of protein tyrosine kinase(s)," J. Pharm. And Experimental Therapeutics, 1993, 267(3), 1119-1125.	
	FU	Schroeder, K. S., et al., "FLIPR: A new instrument for accurate, high throughput optical screening," J. Biomolecular Screening, 1996, 1, 75-80.	
-	FV	Schroeder, K. S., et al., "FLIPR: A new instrument for accurate, high throughput optical screening," J. Biomolecular Screening, 1996, 1, 75-80.	
	FW	Schroeder, K. S., et al., "FLIPR: A new instrument for accurate, high throughput optical screening," J. Biomolecular Screening, 1996, 1, 75-80.	

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	FX	Segal, D. J., et al., "Toward controlling gene expression at will: Selection and design of zine finger domains recognizing each of the 5'-GNN-3' DNA target sequences," Proc. Natl. Acad. Sci. USA, 1999, 96, 2758-2763.	
	FY	Sikora, E., et al., "Quinazoline CB 3717 and CB 3703 inhibition of folate retention and metabolism in ehrlich ascites carcinoma cells and some organs of the host-mouse," Cancer Letters, 1984, 23, 289-295.	
	FZ	Sikora, E., et al., "Development of an assay for the estimation of N.sup.10 -propargyl-5,8-dideazafolic acid polyglutamates in tumor cells," Analytical Biochemistry, 1988, 172, 344-355.	
	GA	Sim, L. J., et al., "Identification of opioid receptor-like (ORL1) peptide-stimulated [.sup.35 S]GTP.gamma.S binding in rat brain," Neuroreport, 1996, 7, 729-733.	
	GB	Smith, T. F., et al., "Comparison of biosequences," Adv. Appl. Math., 1981, 2, 482-489.	
	GC	Smith-Swintosky, V. L., et al., "Protease-activated receptor-2 (PAR-2_is present in the rat hippocampus and is associated with neurodegeneration," J. Neurocham, 1997, 69, 1890-1896.	
	GD	Stables, J., et al., "A bioluminescent assay for agonist activity at potentially any G-protein-coupled receptor," Analytical Biochemistry, 1997, 252, 115-126.	
	GE	Stratowa, C., et al., "Use of a luciferase reporter system for characterizing G-protein-linked receptors," Current Opinion in Biotechnology, 1995, 6, 574-581.	
	GF	Strosberg, et al., "Functional expression of receptors in microorganisms," Trends in Pharmacological Sciences, 1992, 13, 95-98.	
	GG	Strosberg, A. D., et al., "Structure/function relationship of proteins belonging to the family of receptors coupled to GTP-binding proteins,," Eur. J. Biochem., 1991, 196, 1-10.	
	GH	Suidan, H. A., et al., "The thrombin receptor in the nervous system," Semin Thromb Hemost, 1996, 22(2), 125-133.	
	GI	Sutherland, E. W., et al., "Some aspects of the biological role of adenosine 3',5'-monophosphate (cyclic AMP)," Circulation, 1968, 37, 279-306.	

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Examiner	Date	
Signature	Considered	

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( 5	Substitute	for form 1449	В/РТО			Complete if Known		
					ınc	Application Number	10/526,893	
INFORMATION DISCLOSURE				_		Filing Date	Herewith	
•	STATEMENT BY APPLICANT					First Named Inventor	David E. Lowery	
						Art Unit	To Be Determined	
	(Use as many sheets as necessary)					Examiner Name	To Be Determined	
	Sheet 15 of 19					Attorney Docket Number	PHRM0002-105	

		NON PATENT LITERATURE DOCUMENTS	
Examiner Initials *	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T²
	GJ	Sweetnam, P. M., et al., "The role of receptor binding in drug discovery," J. Natural Products, 1993, 56(4), 441-455.	
	GK	Tempest, P. R., et al., "Reshaping a human monoclonal antibody to inhibit human respiratory syncytial virus infection in vivo," Bio/Technology, 1991, 9, 266-271.	
	GL	Torfs, H. et al., "Characterization of a receptor for insect tachykinin-like peptide agonists by functional expression in a stable Drosophila Schneider 2 Cell Line", J. Neurochem., 2000, 74, 2182-2189.	
	GM	Trejo, J., et al., "The cloned thrombin receptor is necessary and sufficient for activation of mitogen-activated protein kinase and mitogenesis in mouse lung fibroblasts," J. Biol. Chem., 1996, 271, 21536-21541.	
	GN	Turgeon, V. L., et al., "Thrombin perturbs neurite outgrowth and induces apoptotic cell death in enriched chick spinal motoneuron cultures through caspase activation," J. Neurosci, 1998, 18(17), 6882-6891.	
	GO	Ubl, J. J., et al., "Characteristics of thrombin-induced calcium signals in rat astrocytes," Glia, 1997, 21, 361-369.	
	GP	Vanden Broeck, "G-protein-coupled receptors in insect cells", Int. Rev. Cytology, 1996, 164, 189-268.	
	GQ	Verhoeyen, M., et al., "reshaping human antibodies: Grafting an antilysozyme activity," Science, 1988, 239, 1534-1536.	
	GR	Voet et al. Biochemistry. 1990. John Wiley & Sons, Inc pp. 126-128 and 228-234.	
	GS	Wieboldt, R., et al., "Immunoaffinity ultrafiltration with ion spray HPLC/MS for screening small-molecule libraries," Anal. Chem., 1997, 69(9), 1683-1691.	
	GT	Williams, M., "Receptor binding in the drug discovery process," Medicinal Research Reviews, 1991, 11(2), 147-184.	

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Signature	Considered	

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Substitute for form 1449B/PTO Complete if Known Application Number 10/526,893 INFORMATION DISCLOSURE Filing Date Herewith STATEMENT BY APPLICANT First Named Inventor David E. Lowery Art Unit To Be Determined (Use as many sheets as necessary) Examiner Name To Be Determined PHRM0002-105 Attorney Docket Number Sheet 16 of

		NON PATENT LITERATURE DOCUMENTS	
Examiner Initials *	Cite No.1	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T²
_	GU	Wolbring, G., et al., "Inhibition of GTP-utilizing enzymes by tyrphostins," J. Biol. Chem., 1994, 269(36), 22470-22472.	
	GV	Wu, H., et al., "Building zinc fingers by selection: toward a therapeutic application," Proc. Natl. Acad. Sci. USA, 1995, 92, 344-348.	
	GW	Yoneda, T., et al., "The antiproliferative effects of tyrosine kinase inhibitors tyrphostins on a human squamous cell carcinoma in vitro and in nude mice," Cancer Research, 1991, 51, 4430-4435.	
	GX	Adams, M.D., et al., "The genome sequence of drosophila melanogaster," EMBL/GenBank/DDBJ, XP-002176201, Mar. 21, 2000, 3 pages.	
	GY	Alcedo, J., et al., "The drosophila smoothened gene encodes a seven-pass membrane protein, a putative receptor for the hedgehog signal," Cell, XP-002166694, Jul. 26, 1996, 86, 221-232.	
	GZ	Celniker, S.E., et al., "Drosophila melanogaster, chromosome X, region 17C-17E," EMBL, XP-002176202, Oct. 22, 1999, 2 pages.	
	НА	Celniker, S.E., et al., "Drosophila melanogaster, chromosome 2R, region 42A8-42A16, P1 clones DS06954 and DS05325," EMBL, XP-002176200, Mar. 24, 1999, 2 pages.	
	НВ	Celniker, S.E., et al., "Drosophila melanogaster, chromosome 3R, region 83D-83D, BAC clone BACR26C09," EMBL, XP-002176198, Sep. 17, 1999, 2 pages.	
	НС	Muzny, D.M., et al., "Drosophila melanogaster clone RPC198-10L1," EMBL, XP-002166695, Aug. 23, 1999, 3 pages.	
	HD	Muzny, D.M., et al., "Drosophila melanogaster clone RPCI98-23M20," EMBL, xP-002176199, Aug. 23, 1999, 3 pages.	
	HE	Nichols, R., "Isolation and structural characterization of drosophila TDVDHVFLRF amide and FMRF amide-containing neural peptides," Medline, XP-002166696, 1992, 1 page.	

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Substitute for form 1449B/PTO				Complete if Known		
				Application Number	10/526,893	
			CLOSURE	Filing Date	Herewith	
STAT	EMENT	BY AF	PPLICANT	First Named Inventor	David E. Lowery	
				Art Unit	To Be Determined	
	(Use as many	sheets as	necessary)	Examiner Name	To Be Determined	
Sheet	17	of	19	Attorney Docket Number	PHRM0002-105	

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	HF	Taghert, P.H., et al., "Interspecific comparison of a drosophila gene encoding FMRF amiderelated neuropeptides," J. Neuroscience, USA,, 1990, 10(6), 1929-1942.	
	HG	Copy of PCT International Search Report dated Oct. 29, 2001 for International Application No. PCT/US00/29002.	
	*HI	Berger et al., "Guide to Molecular Cloning Techniques," Methods in Enzymology, Academic Press, Inc., San Diego, CA 1987.	
	*HI	Cobbold et al., "Aequorin measurements of cytoplasmic free calcium," McCormack J.G., et al. (Eds.), Cellular Calcium: A Practial Approach (1991) Oxford, IRL Press.	
	*НЈ	Current Protocols in Molecular Biology, John Wiley & Sons, NY 1999.	
	*HK	Eisenthal et al., Enzyme Assays: A Practical Approach, Oxford University Press, 1992.	
	*HL	Harlow et al., Antibodies: A Laboratory Manual, Cold Spring Harbor Laboratory, Cold Spring Harbor, NY, 1988.	
	*HM	Haugland, Handbook of Fluorescent Probes and Research Chemicals, 6th Ed., 1996, Eugene OR: Molecular Probes.	
	*HN	Hendix, (ed.), Lambda II, Cold Spring Harbor Press, Cold Spring Harbor, NY 1980.	
<del></del>	*НО	Hershey (ed.), The Bacteriophage Lambda, Cold Spring Harbor Press, Cold Spring Harbor, NY, 1973.	
	*HP	Kruse et al. (eds). Tissue Culture, Academic Press, 1973.	

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			CLOSURE	Filing Date	Herewith
STAT	EMENT	BY A	PPLICANT	First Named Inventor	David E. Lowery
				Art Unit	To Be Determined
	(Use as many	sheets as	necessary)	Examiner Name	To Be Determined
Sheet	18	of	19	Attorney Docket Number	PHRM0002-105

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	HQ	O'Rielly et al. (eds.), Baculovirus Expression Vectors: A Laboratory Manual, W.H. Freeman and Company, New York, 1992.	
	HR	Sambrook et al., Molecular Cloning: A Laboratory Manual, second edition, Cold Spring Harbor Press, Cold Spring Harbor, NY 1989.	,
	HS	Stapleton et al., "A Drosophila full-length cDNA Resource," Genome Biology (2002) 3(12):1-8.	
	нт	International Search Report dated April 19, 2004 for International Application No. PCT/US03/24488.	
	HU	Garcynski, et al., "Characterization of a functional neuropeptide F receptor from Drosophila melanogaster," Peptides (2002) 23:773-780	
	HV	Holmes, et al., "Cloning and transcriptional expression of leucokinin-like peptide receptor from the Southern cattle tick, Boophilus microplus (Acari:Ixodidae)," Insect Mol. Biol. (2000) 9:457-465	
	HW	Birgul, et al., "Reverse physiology in Drosophila: identification of a novel allatostatin-like neuropeptide and its cognate receptor structurally related to the mammalian somatostatin/galanin/opiod receptor family," EMBO J. 18:5892-5900	
	нх	Cazzamali, et al., "Molecular cloning and functional expression of a Drosophila corazonin receptor," Biochem. Biophys. Res. Comm. (2002) 298:31-36	
	НҮ	Larsen, et al., "Type A allatostatinsfrom Drosophila melanogaster and Diplotera punctata activate two Drosophila allatostatin receptors, DAR-1 and DAR-2, expressed in CHO cells," Biochem. Biophys. Res. Comm. (2001) 286:895-901	
	HZ	Nichols, "Isolation and expression of the Drosophila drosufakinin neural peptide gene product DSK-I," Mol. Cell. Neurosci. (1992) 3:342-347	
	IA	O'Donnell, et al., "Hormonally controlled chloride movement across Drosophila tubules is via ion channels in stellate cells," Am. J. Physiol. (1998) 43:R1039-R1049	

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Signature	 Considered	

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Substitute	for form 1449B/P	то		Complete if Known		
=0			01 001105	Application Number	10/526,893	
			CLOSURE	Filing Date	Herewith	
STAT	STATEMENT BY APPLICANT			First Named Inventor	David E. Lowery	
				Art Unit	To Be Determined	
(Use as many sheets as necessary)			necessary)	Examiner Name	To Be Determined	
Sheet	19	of	19	Attorney Docket Number	PHRM0002-105	

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	IB	Mertens, et al., "Characterization of the short neuropeptide F receptor from Drosophila melanogaster," Biochem. Biophys. Res. Comm. (2002) 1140-1148					
· ·	IC	Siviter, et al., "Expression and functional characterization of a Drosophila neuropeptide precursor with homology to mammalian preprotachykinin A," J. Biol. Chem. (2000) 275:23273-23280					
	ID	Staubli, et al., "Molecular identification of the insect adipokinetic hormone receptors," Proc. Natl. Acad. Sci. USA (2002) 99:3446-3451					
	IE	Price et al., "Drosophila melanogaster flatline encodes a myotropin orthologue to Manduca sexta allatostatin," Peptides (2002) 23:787-794					
	IF	Kubiak, et al., "Cloning and Functional Expression of the first Drosophila melanogaster sulfakinin receptor DSK-R1," Biochem. Biophys. Res. Comm. (2002) 291:313-320					
	IG	Radford, et al., "Systematic G-protein-coupled receptor analysis in Drosophila melanogaster identifies a leucokinin receptor with novel roles," J. Biol. Chem. (2002) 277:38810-38817					
	IH	Williamson, et al, "Molecular cloning, genomic organization, and expression of a C-type (Manduca sexta-type) allatostatin preprohormone from Drosophila melanogaster," Biochem. Biophys. Res. Comm. (2001) 282:124-130	- - - -				
	П	Cazzamali, et al., "Molecular cloning and functional expression of the first insect FMRFamide receptor," Proc. natl. Acad. Sci. USA (2002) 99:12073-12078					
	IJ	Kreikenkamp et al., "Functional annotation of two orphan G-protein-coupled receptors, drostar1 and -2, from Drosophila melanogaster and their ligands by reverse pharmacology," J. Biol. Chem. (2002) 277:39937-39943					
	IK	Park et al., "Identification of G protein-coupled receptors for Drosophila PRXamide peptides, CCAP, corazonin, and AKH supports a theory of ligand-receptor coevolution," Proc. Natl. Acad. Sci. USA (2002) 99:11423-11428					

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Signature	Considered	

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